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Commissioner for Patents

Alexandria, VA 22313-1450

P.O. Box 1450

TOWNSEND and TOWNSEND and CREW LLP

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re application of:

Thomas Cech et al.

Application No.: 10/044,692

Filed: January 11, 2002

For: HUMAN TELOMERASE

CATALYTIC SUBUNIT: DIAGNOSTIC

AND THERAPEUTIC METHODS

Examiner:

Art Unit:

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT UNDER 37

Attorney Docket No.: 015389-00

Client Reference No.: 018/213C

CFR §1.97 and §1.98

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

Sir:

The references cited on attached form PTO/SB/08A and PTO/SB/08B are being called to the attention of the Examiner. Copies of the references are enclosed. It is respectfully requested that the cited references be expressly considered during the prosecution of this application, and the references be made of record therein and appear among the "references cited" on any patent to issue therefrom.

As provided for by 37 CFR 1.97(g) and (h), no inference should be made that the information and references cited are prior art merely because they are in this statement and no

representation is being made that a search has been conducted or that this statement encompasses all the possible relevant information.

Applicant believes that <u>no fee is required</u> for submission of this statement. However, if a fee is required, the Commissioner is authorized to deduct such fee from the undersigned's Deposit Account No. 20-1430. Please deduct any additional fees from, or credit any overpayment to, the above-noted Deposit Account.

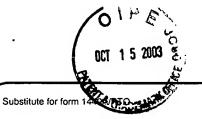
Respectfully submitted,

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## INFORMATION DISCLOSURE STATEMENT BY APPLICANT

(use as many sheets as necessary)

Sheet	1	of	5

	Complete if Known
Application Number	10/044,692
Filing Date	January 11, 2002
First Named Inventor	Cech
Art Unit	
Examiner Name	
Attorney Docket Number	015389-002640US

	U.S. PATENT DOCUMENTS+					
		Document Number				
Examiner Initials*	Cite No. <sup>1</sup>	Number Kind Code <sup>2</sup> (if known)	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	
	A1	US-5,597,697	01-28-1997	Diamond		
	A2	US-5,917,025	06-29-1999	Collins		
	А3	US-6,166,178	12-26-2000	Cech et al.		
	A4	US-6,309,867	10-30-2001	Cech et al.		
	A5	US-6,337,200	01-08-2002	Morin		
	A6	US-6,475,789 B1	11-05-2002	Cech et al.		
	A7	US-6,610,839 B1	08-26-2003	Cech et al.		
	A8	US-6,617,110 B1	09-09-2003	Cech et al.		

	FOREIGN PATENT DOCUMENTS							
Examiner Initials*	Cite No. <sup>1</sup>	For Country Code <sup>3</sup>	eign Patent Docume Number <sup>4</sup> Kind	nt 1 Code <sup>5</sup> ( <i>if known</i> )	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T <sup>6</sup>
	B1	GB	2 317 891	Α	04-08-1998			
	B2	PCT	WO 93/23572	A1	11-25-1993			
	В3	PCT	WO 94/17210	A1	08-04-1994			
	B4	PCT	WO 98/14592	A2	04-09-1998			
	B5	PCT	WO 98/14593	A2	04-09-1998			
	В6	PCT	WO 98/23759	A2	06-04-1998			
	B7	PCT	WO 99/33998	A2	07-08-1999			
	B8	PCT	WO 99/38964	A2	08-05-1999			
	B9	PCT	WO 00/46355	A2	08-10-2000			

Examiner Signature	 Date Considered	

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## **INFORMATION DISCLOSURE** STATEMENT BY APPLICANT

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Sheet 2 of 5

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		NON PATENT LITERATURE DOCUMENTS	
Examiner Initials *	Cite No.1	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T 2
	C1	ADAMSON, D. et al. "Significant Telomere Shortening in Childhood Leukemia", Cancer Genet. Cytogenet, 1992; pp. 204-206, Vol. 61	
	C2	AUTEXIER, CHANTAL, et al., "Telomerase and cancer: revisiting the telomere hypothesis; Trends in Biochemical Sciences, 1996, pp. 387-391, Vol. 10, No. 21.	
	СЗ	AVILION, A., "Characterization and expression of human telomerase," Dissertation Abstracts International", 1996, pp. 5930-B, Vol. 56, No. 11	
	C4	BARINGA, MARCIA, "The Telomerase Picture Fills In", Science, April 25, 1997; pp. 528-529, Vol. 276	
	C5	CHIU, et al. "Replicative senescence and cell immortality: the role of telomeres and telomerase (44075)", <i>Proc. Soc. Exp. Bio. Med.</i> , 1997, pp. 99-106, Vol. 214.	
	C6	CHONG, L. et al. "A Human Telomeric Protein", Science, Dec. 1995, pp. 1663-1667, Vol. 270.	
	C7	COLLINS, KATHLEEN, "Structure and function of telomerase", Current Opinion in Cell Biology, 1996, pp. 374-380, Vol. 8	
	C8	COUNTER, C. et al. "Telomerase Activity in Normal Leukocytes and in Hematologic Malignancies" <i>Blood</i> ; May 1, 1995; pp. 2315-2320, Vol. 85, No. 9.	
	C9	COUNTER, C. et al. "Telomere shortening associated with chromosome instability is arrested in immortal cells which express telomerase activity", <i>The EMBO Journal</i> ; 1992, pp. 1921-1929, Vol. 11; No. 5, Oxford University Press.	
	C10	DE LANGE, T. et al. "Structure and Variability of Human Chromosome Ends"; <i>Molecular and Cellular Biology;</i> February 1990; pp. 518-527, Vol. 10, No. 2.	
	C11	FLAVELL, R. & MATHIAS, R. "Prospects for transforming monocot crop plants", <i>Nature</i> , 12 Jan. 1984, pp. 108-109, Vol. 307.	
	C12	FRESHNEY, <u>Culture of Animal Cells, A Manual Basic Technique</u> , p. 4, Alan R. Liss, Inc. New York, 1983.	
	C13	Genbank Accession No. A46242; September 21, 1993	
	C14	Genbank Accession No. L38903; January 30, 1995	
	C15	Genbank Accession No. Q06163; November 01, 1995	
	C16	Genbank Accession No. S39696; October 07, 1994	

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PTO/SB/08B (08-03) stitute for form 1449B/PTO Complete if Known Application Number 10/044,692 INFORMATION DISCLOSURE Filing Date January 11, 2002 STATEMENT BY APPLICANT First Named Inventor Cech Art Unit (use as many sheets as necessary) Examiner Name Sheet of 5 Attorney Docket Number 015389-002640US

	NON PATENT LITERATURE DOCUMENTS						
Examiner Initials *	Cite No.1	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T²				
	C17	Genbank Accession No. S53396; May 5, 1995					
	C18	GenBank Accession No. U95964; May 5, 1997					
	C19	GenBank Accession No. W70315; June 19, 1996					
	C20	GLASER, P. et al. "Bacillus subtilis genome project: cloning and sequencing of the 97 kb region from 325° to 333°" <i>Molecular Microbiology</i> , 1993, pp. 371-384, Vol. 10, No. 2.					
	C21	GOODMAN, R. et al. "Gene Transfer in Crop Improvement", Science, 3 April 1997, pp. 48-54, Vol. 236.					
	C22	GREIDER, C. "Telomeres, Telomerase and Senescence"; <i>BioEssays</i> ; 1990; pp. 363-369, Vol. 12, No. 8.					
	C23	HARLEY, C. "Telomere loss: Mitotic clock or genetic time bomb?" <i>Mutation Research</i> ; 1991; pp. 271-282, Vol. 256, Elsevier Science Publishers.					
	C24	HARLEY, C. & VILLEPONTEAU, B. "Telomeres and telomerase in aging and cancer" Current Opinion in Genetics and Development; 1995, pp. 249-255, Vol. 5.					
	C25	HARLEY, C. et al. "Telomeres shorten during ageing of human fibroblasts", <i>Nature</i> , 31 May 1990; pp. 458-460, Vol. 345.					
	C26	HASTIE, N. et al. "Telomere reduction in human colorectal carcinoma and with ageing", <i>Nature</i> , 30 August 1990, pp. 866-868, Vol. 346.					
	C27	HEALY, K. C. "Telomere dynamics and telomerase activation in tumor progression: prospects for prognosis and therapy" <i>Oncol. Res.</i> , 1995, pp. 121-130, Vol. 7					
	C28	HENDERSON, C. Cancer genetics gene regulates telomerase resulting in death of cancer cells; Gene Therapy Weekly; 11 September 1995.					
	C29	HIYAMA, E. et. al. "Correlating telomerase activity levels with human neuroblastoma outcomes"; Nature Medicine; 3 March 1995; pp. 249-255, Vol. 1, No. 3.					
	C30	HOLTZMANN, K. et al. "Telomeric Associations and Loss of Telomeric DNA Repeats in Renal Tumors", Genes, Chromosomes & Cancer, 1993, pp. 178-181, Vol. 6.					
	C31	JÄHNE, A. et al. "Genetic Engineering of Cereal Crop Plants: A Review", Euphytica, 1995, pp. 35-44, Vol. 85, Kluwer Academic Publishers, Netherlands.					
	C32	JOLLIFFE, L.K. "Humanized antibodies: enhancing therapeutic utility through antibody engineering", Int. Rev. Immunol., 1993, pp. 241-250, Vol. 10					
	C33	KIM, N. et al. "Specific Association of Human Telomerase Activity with Immortal Cells and Cancer", <i>Science</i> , 23 December 1994, pp. 2011-2014, Vol. 266.					
Examiner Signature		Date Considered					

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<del></del>	T	NON PATENT LITERATURE DOCUMENTS	
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	C34	KLINGELHUTZ, A. et al. "Restoration of Telomeres in Human Papillomavirus- Immortalized Human Anogenital Epithelial Cells"; <i>Molecular and Cellular Biology</i> ; Feb. 1994, pp. 961-969, Vol. 14, No. 2.	
	C35	LEWIS, A. & CROWE, J.S. "Generation of humanized monoclonal antibodies by 'best fit' framework selection and recombinant polymerase chain reaction", <i>Year Immunol.</i> , 1993, pp. 110-118, Vol. 7	
	C36	Linking telomerase and tumors; <i>Genesis Report- Dx</i> ; 1995; Vol. 4, No. 6; Publisher Genesis Group Associates.	
	C37	LUNDBLAD, V. & BLACKBURN, E., Letter to the Editor entitled, "RNA-Dependent Polymerase Motifs in EST1: Tentative Identification of a Protein Component of an Essential Yeast Telomerase", Cell, 23 Feb. 1990, pp. 529-530, Vol. 60.	
	C38	LUSTIG, ARTHUR J., "The identification of telomerase subunits: catalysing telomere research", Trends in Cell Biology; August 1997, pp. 299-302, Vol. 7.	
	C39	MALICKI, J. et al. "A human <i>HOX4B</i> regulatory element provides head-specific expression in <i>Drosophila</i> embryos", Nature, 23 July 1992, pp. 345-357, Vol. 358.	
	C40	NAKAYAMA, J. et al. "Cloning of a Candidate cDNA Encoding a Proteinaceous Component of Mammalian Telomerase", <i>Molecular Biology Cell Abstracts Supp.</i> 7, 1996, p. 875-884, 286a Section 1664.	
	C41	NATARAJAN et al. "Major histocompatibility complex determinants select T-cell receptor alpha chain variable region dominance in a peptide-specific response." <i>Proc. Natl. Acad. Sci.</i> , Oct 1992, pp. 8874-8878, Vol. 19.	
	C42	PASZKOWSKI, JERZY et al. "Direct gene transfer to plants," The EMBO Journal, 1984, pp. 2717-2722, Vol. 3, No. 12.	
	C43	POTRYKUS, I. et al. "Direct gene transfer to cells of a graminaceous monocot", <i>Mol. Gen. Genet.</i> , 1985, pp. 183-188, Vol. 199.	
	C44 RAYMOND, E., et al.; Agents that target telomerase and telomeres; <i>Curr. Opin. Biotechnol.</i> ; 1996; 7:583-91		
	C45	RHYU, M.S. "Telomeres, telomerase, and immortality"; J. Natl. Cancer Inst.; 21 June 1995; pp. 884-894, Vol. 87, No. 12.	
	C46	SCHENA et al. "Parallel human genome analysis: Microarray-based expression monitoring of 1000 genes", <i>Proc. Natl. Acad. Sci.</i> , Oct. 1996, pp. 10614-10619, Vol. 93, USA.	
	C47	SCHWARTZ, H. et al. "Telomere Reduction in Giant Cell Tumor of Bone and with Aging"; Cancer Genet Cytogenet; 1993; pp. 132-138, Vol. 71, Elsevier Science Publishing Co., Inc., New York, U.S.A.	
	C48	SINGER, M. "Unusual Reverse Transcriptases", Journal of Biological Chemistry; 1995, pp. 24623-23626, Vol. 270, No. 42	
	C49	SMITH, J. & YEH, G. "Telomere Reduction in Endometrial Adenocarcinoma"; <i>Am. J. Obstet. Gynecol.</i> ; Dec. 1992; pp. 1883-1887, Vol. 167, No. 6	

Signature Considered

EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

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Sheet	5	of	5

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NON PATENT LITERATURE DOCUMENTS							
Examiner Initials *	(2000), value page(b), value location						
	C50	TAIT, J. et al. "Structure and Polymorphisms of the Human Annexin III (ANX3) Gene", <i>Genomics</i> , 1993, pp. 79-86, Vol. 18, No. 1					
	C51	WINTER, G. & HARRIS, W. "Humanized Antibodies", <i>Trends Pharmacol. Sci.</i> , May 1993, pp. 139-143, Vol. 14.					
	C52	WIRTH, URS et al; "Immediate-Early RNA 2.9 and Early RNA 2.6 of Bovine Herpesvirus 1 Are 3' Coterminal and Encode of Putative Zinc Finger Transactivator Protein"; <i>Journal of Virology</i> ; May 1992; pp. 2763-2772, Vol. 66, No. 5.					
	C53	ZAUG, ARTHUR et al., "Method for determining RNA 3' ends and application to human telomerase RNA", <i>Nucleic Acids Research</i> , 1996, pp. 532-533, Vol. 24, No. 3.					

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